STUDIES IN THE TREATMENT OF MALARIA

XXI. ARSENIC IN SIMPLE TERTIAN MALARIA

BY

LIEUT.-COL. J. W. W. STEPHENS, R.A.M.C.
W. YORKE
B. BLACKLOCK
J. W. S. MACFIE
C. FORSTER COOPER
AND

From the Liverpool School of Tropical Medicine

H. F. CARTER

Undertaken at the request of the War Office

(Received for publication 3 December, 1918)

Many statements concerning the value of arsenic in the treatment of malaria are found in the literature, but with the exception of a few more recent records relating to organic preparations we have been unable to discover the evidence on which such statements are based.

We proceeded therefore to determine (1) whether inorganic arsenic alone has any action—palliative or curative—in simple tertian malaria, and if so in what dose it should be administered; and (2) whether a combination of arsenic and quinine is more effective than quinine alone. The observations made by us can be divided into the following four groups:—

GROUP I. Small doses of arsenic.

A. Liquor arsenicalis* minims 15 daily, for 8 weeks.

B. The same as Treatment A, with an injection of Quinine 2 HCl grains 15 intramuscularly on each of the first two days only.

^{*} The British Pharmacopocia preparation containing 1 per cent. of arsenious anhydride.

GROUP II. Small doses of arsenic, or of arsenic and strychnine, with quinine.

- C. Liquor arsenici HCl* minims 15 + Quinine HCl grains 5, daily for 8 weeks.
- D. Liquor arsenici HCl minims 15 and Liquor strychninae HCl minims 15 + Quinine HCl grains 5, daily for 8 weeks
- E. Control Series. Quinine HCl grains 5 daily for 8 weeks.

GROUP III. Large doses of arsenic.

- F. Liquor arsenicalis minims 30 daily, with one or two periods of intermission, for 8 weeks.
- G. The same as Treatment F, with an injection of Quinine 2 HCl grains 15 intramuscularly on each of the first two days only.
- H. Control series. Quinine 2 HCl grains 15 intramuscularly on each of two consecutive days only.

All the cases were adult males, the great majority of whom were infected either in Macedonia or in East Africa (Tables XIV to XIX). In every instance a diagnosis of simple tertian malaria was made microscopically, and in all cases parasites were present in the blood on the day treatment commenced. Blood examinations were made daily.

The records of the observations are given in the tables and charts. In the tables:—

o = absence of fever and parasites.

1, 2, etc. = number of parasitic febrile relapses, weekly.

1*, 2*, etc. = number of non-parasitic febrile attacks, weekly.

P¹, P², etc. = number of days on which parasites were present weekly.

Remarks = post-treatment observations.

In the charts:—

T. = simple tertian trophozoites or schizonts.

G. = simple tertian gametes.

Neg. = no parasites found.

Note.— Λ rise of temperature above 100° F., of which the nature is unknown, is termed a *febrile attack*. A similar rise of temperature accompanied by parasites in the blood at the time, or within three days, is termed a *parasitic febrile relapse* or *true relapse*. The term *paroxysm* is used indifferently to denote any febrile disturbance of 100° F. or more.

As we have pointed out elsewhere, the effect of any treatment may be considered from two points of view: (1) the *palliative* action, i.e., the degree to which symptoms are controlled, and the blood kept free

^{*} The British Pharmacopoeia preparation containing 1 per cent. of arsenious anhydride.

from parasites during the treatment; and (2) the curative action, i.e., whether or no relapses occur during the observation period* after cessation of treatment. In order that the palliative results obtained in the various series of observations may have a comparative value, it is necessary to express the number of cases having true relapses, of those having febrile attacks, and of those having parasitic relapses, as percentages of the total cases undergoing treatment in any particular week. In Tables II, IV, VI, IX and XI, the following sets of figures, each having a comparative value, are given:—

- 1. The number of cases which had parasitic febrile relapses each week, expressed as a percentage of all cases treated.
- 2. The number of parasitic febrile relapses experienced per week by each parasitic febrile relapse case.
- 3. The number of cases which had febrile paroxysms (parasitic and non-parasitic) each week, expressed as a percentage of all cases treated.
- 4. The number of febrile paroxysms (parasitic and non-parasitic) experienced per week by each febrile (parasitic and non-parasitic) case.
- 5. The number of cases which had parasitic relapses (febrile and non-febrile) each week, expressed as a percentage of all cases treated.
- 6. The number of parasitic relapses (febrile and non-febrile) experienced per week by each parasitic relapse (febrile and non-febrile) case.

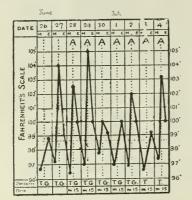
GROUP I. Small doses of arsenic

A. Liquor arsenicalis minims 15 daily.† (Cases 1186 to 1188). Three cases only were treated in this manner, as it was soon found that Liquor arsenicalis in this dose failed to cause the disappearance of parasites from the cutaneous blood or to control the fever. In two cases (1186 and 1187) it was found necessary to alter the treatment after seven days, in the third case (1188) after twenty-two days. Details are given in Charts 1186 to 1188.

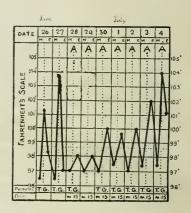
^{*} This, as in all our previous papers, is 60 days—an entirely arbitrary period.

[†] Liq. arsenicalis M v Aq. chloroformi ad 3 i 3 i. t.d.s.

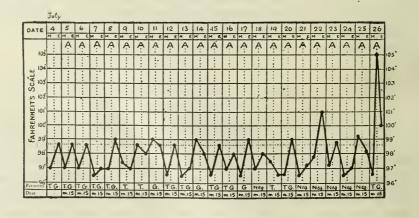
Case 1186



Case 1187



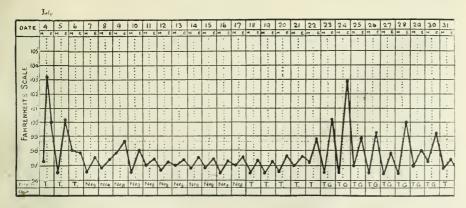
CASE 1188



B. The same as Treatment A, with an injection of Quinine 2 HCl grains 15 intramuscularly on each of the first two days only (Cases 1189 and 1190).

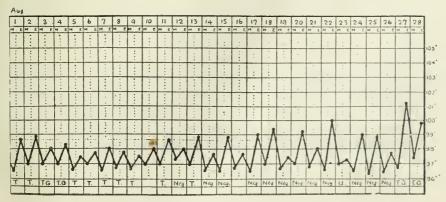
The object of this series of observations was to determine whether, having caused the disappearance of parasites from the cutaneous blood and having controlled symptoms by two initial intramuscular injections of quinine, the daily administration of Liquor arsenicalis (minims 15) would suffice to prevent relapses (parasitic or febrile). The two cases treated in this way relapsed on the thirteenth and eighteenth days respectively after the commencement of treatment, and subsequently parasites were found more or less frequently during the course of treatment; consequently it was considered unnecessary to subject further cases to this form of treatment. Details are given in Charts 1189 and 1190.

Case 1189 (Part I)

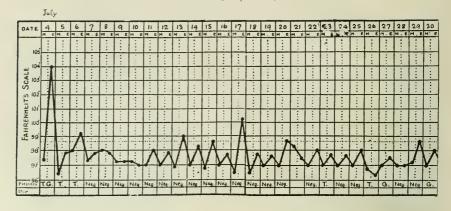


Treatment commenced 5 July.

Case 1189 (Part II)

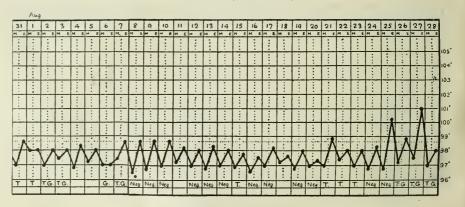


CASE 1190 (Part I)



Treatment commenced 5 July.

CASE 1190 (Part II)



CONCLUSION

Liquor arsenicalis alone, minims 15 daily, fails to cause the disappearance of parasites or to control the symptoms, and does not suffice to prevent relapses (parasitic and febrile) in cases in which parasites and fever have disappeared as the result of two initial intramuscular injections of quinine bihydrochloride.

GROUP II. Small doses of arsenic, or of arsenic and strychnine, with quinine

We next proceeded to ascertain whether combinations of (1) arsenic + quinine, or (2) arsenic and strychnine (on account of the alleged tonic effect of the latter) + quinine, were more effective than quinine alone. For this purpose the following three series (C, D, and E) of observations were undertaken.

C. Liquor arsenici HCl minims 15 + Quinine HCl grains 5 daily* for eight weeks (Cases 1191 to 1218).

In two of the twenty-eight cases treatment was commenced during an apyrexial period; in the remaining twenty-six cases the temperature fell to normal† within five days of the beginning of treatment.

In twenty-one cases parasites disappeared from the cutaneous blood in one to seven days. In the remaining seven (Nos. 1194, 1197, 1206, 1209, 1210, 1214 and 1218) parasites persisted practically throughout treatment (Table XIV).

RELAPSES

During treatment. In five of the twenty-eight cases, owing to the severity of the relapses and the grave clinical condition of the patients, it was found impossible to continue the treatment for the full period of eight weeks, e.g., in Cases 1214 and 1215 treatment had to be changed in the eighth week, in Case 1216 in the seventh week, and in Cases 1217 and 1218 in the fifth week. Consequently the number of cases under treatment was in the first week twenty-eight and in the eighth week twenty-three. The average weekly number, over a period of eight weeks, of cases which had (1) parasitic febrile relapses was 25'7 per cent of cases treated, (2) non-parasitic febrile attacks 6'3 per cent., (3) febrile paroxysms (parasitic and non-parasitic) 32'0 per cent., (4) parasitic relapses (febrile and non-febrile) 42'4 per cent. (Tables I and II).

After treatment. Eighteen of the twenty-three cases observed after cessation of treatment relapsed within the sixty days' observation period. Parasites re-appeared in one to thirty-seven days and

^{*} Liq. arsenici HCl M v Quininae HCl gr. 18 Aq. chloroformi ad 5 i 3i. t.d.s.

[†] By this is meant that the temperature fell to normal and remained so for at least two days.

TABLE I.

C. Summary of results of oral administration of Liquor arsenici HCl minims 15 + Quinine HCl grains 5, daily for 8 weeks, in simple tertian malaria.

Week of Treatment	ıst	2nd	3rd	4th	5th	6th	7th	8th
Number of cases treated	28	28	28	28	26	26	25	23
Number of cases having parasitic febrile relapses	I	6	5	6	5	7	11	5
Number of cases having non-parasitic febrile attacks	1	0	4	5	2	4	1	+
Crand total of all febrile cases	2	6	9	11	7	11	12	9
Total number of parasitic febrile relapses	1	11	8	14	9	18	23	12
Total number of non-parasitic febrile attacks	1	0	+	8	2	4	1	8
Crand total of all febrile par- oxysms	2	11	12	22	11	22	24	20
Number of parasitic cases (febrile and non-febrile)	14	8	9	1.1	-9	11	15	12
Total number of occasions parasites found	28	44	47	48	39	43	60	42

TABLE II.
Analysis of TABLE I.

Week of Treatment	ıst	2nd	3rd	4th	5th	6th	7th	8th	Average per week
Percentage of parasitic febrile relapse cases per cases treated	3.6	21.4	17.8	21.4_	19.2	26.8	44.0	21.7	25•7
Number of parasitic febrile relapses per parasitic febrile relapse case	2.0	1.0	1.8	1.8	1.4	1.6	1.1	1.8	1•5
Percentage of all febrile (parasitic and non- parasitic) cases per cases treated	7.1	21.4	32.1	39.2	26.8	42.3	48.0	39.1	32-0
Number of all febrile (parasitic and non- parasitic) paroxysms per febrile case	1.0	1.8	1.3	2.0	1.6	2.0	2.0	2.2	1•7
Percentage of all parasitic (febrile and non- febrile) cases per cases treated	50.0	28.6	32.1	39.3	34.6	42.3	60.0	52.2	42 • 4
Number of occasions parasites found per parasitic case	2.0	5.5	5.2	4.4	4.3	3.9	4.0	3.2	4.1

febrile relapses occurred in one to thirty-seven days after cessation of treatment. In five of the twenty-eight cases (Nos. 1214 to 1218) the full course of treatment was not completed as the condition was uncontrolled. These therefore should be added to the cases that relapsed, making the total number of failures twenty-three (82 per cent.).

In one of the five cases which did not relapse the post-treatment observation period was less than sixty days, viz., in Case 1195, thirty-seven days. Consequently the minimum number of relapses (those actually observed) was twenty-three and the possible maximum number (those actually observed + the number of cases not relapsing but lost sight of before expiration of the sixty-day post-treatment observation period) was twenty-four, giving a minimum percentage of 82 and a maximum percentage of 86 (Tables XIII and XIV).

D. Liquor arsenici HCl minims 15 and Liquor strychninae HCl minims 15 + Quinine HCl grains 5 daily* for eight weeks (Cases 1219 to 1243).

In four of the twenty-five cases treatment was commenced during an apyrexial period; in the remaining twenty-one cases the temperature fell to normal within eight days of the beginning of treatment.

In twenty-one cases parasites disappeared from the cutaneous blood in one to seven days. In seven cases (1221, 1222, 1238, 1239, 1241, 1242, and 1243) parasites persisted practically throughout treatment (Table XV).

RELAPSES

During treatment. In one (1243) of the twenty-five cases, owing to the severity of the relapses and the grave clinical condition of the patient, it was found necessary to alter the treatment in the sixth week. Consequently the number of cases under treatment was in the first week twenty-five and in the eighth week twenty-four. The average weekly number, over a period of eight weeks, of cases which had (1) parasitic febrile relapses was 13'2 per cent. of cases

^{*} Liq. arsenici HCl M 5 Liq. strychninae HCl M 5 Quininae HCl gr. 1²/₃ Aq. chloroformi ad 3 i 5 i. t.d.s.

treated, (2) non-parasitic febrile attacks 7⁻I per cent., (3) febrile paroxysms (parasitic and non-parasitic) 20⁻3 per cent., and (4) parasitic relapses (febrile and non-febrile) 37⁻6 per cent. (Tables III and IV).

After treatment. Of the twenty-three cases observed after cessation of treatment seventeen relapsed within the sixty-day observation period. Parasites re-appeared in one to twenty-six days and febrile relapses in seven to twenty-seven days after cessation of treatment. In one (1243) of the twenty-five cases the full course of treatment was not completed as the condition was uncontrolled. This case therefore should be added to the cases that relapsed, making the total number of failures eighteen (75 per cent.).

In two of the six cases which did not relapse the post-treatment observation period was less than sixty days, viz., in Case 1223, forty-two days and in Case 1227, thirty-eight days. Consequently the minimum number of relapses (those actually observed) was eighteen, and the possible maximum number (those actually observed + the number of cases not relapsing but lost sight of before expiration of the full post-treatment observation period of sixty days) was twenty, giving a minimum percentage of 75 and a maximum of 83 (Tables XIII and XV).

E. Control series. Quinine HCl grains 5 daily* for eight weeks (Cases 1244 to 1269).

In these twenty-six cases the temperature fell to normal within three days of the beginning of treatment.

In eighteen cases parasites disappeared from the cutaneous blood in from one to ten days. In ten cases (Nos. 1244, 1245, 1248, 1249, 1252, 1258, 1261, 1262, 1266 and 1268) parasites persisted practically throughout treatment (Table XVI).

RELAPSES

During treatment. In one (1268) of the twenty-six cases, as the condition was uncontrolled, the treatment was changed in the

^{*} Quininae HCl gr. 13 Aq. chloroformi ad $\frac{7}{5}$ i $\frac{7}{5}$ i. t.d.s.

TABLE III.

D. Summary of results of oral administration of Liquor arsenici HCl minims 15 and Liquor strychninae HCl minims 15 + Quinine HCl grains 5, daily for 8 weeks, in simple tertian malaria.

Week of Treatment	ıst	2nd	3rd	4th	5th	6th	7th	8th
Number of cases treated	25	25	25	25	25	24	24	24
Number of cases having parasitic febrile relapses	5	3	2	3	6	1	I	2
Number of cases having non-parasitic febrile attacks	1	3	2	1	1	2	ı	3
Grand total of all febrile cases	6	6	4	4	7	6	2	5
Total number of parasitic febrile relapses	10	5	2	6	1.4	4	1	4
Total number of non-parasitic febrile attacks	I	+	2	I	2	3	4	9
Crand total of all febrile par- oxysms	11	9	4	7	16	7	5	13
Number of parasitic cases (febrile and non-febrile)	11	10	9	9	8	8	10	9
Total number of occasions parasites found	2.1	35	31	33	34	26	35	43

Table IV.

Analysis of Table III.

Week of Treatment	tst	2nd	3rd	4th	5th	6th	7th	Sth	Average per week
*	20.0	12.0	8.0	12.0	24.0	16.7	4.2	8.3	13.2
Number of parasitic febrile relapses per parasitic febrile relapse case	2.0	1.7	1.0	2.0	2.3	1.0	1.0	2.0	1.6
Percentage of all febrile (parasitic and non- parasitic) cases per cases treated	24.0	24.0	16.0	16.0	28.0	25.0	8.3	20.8	20.3
Number of all febrile (parasitic and non- parasitic) paroxysms per febrile case	1.8	1.5	1.0	1.7	2.3	1.2	2.5	2.6	1.8
Percentage of all parasitic (febrile and non-febrile) cases per cases treated	11.0	40.0	36.0	36.0	32.0	33.3	41.7	37:5	37.6
Number of occasions parasites found per parasitic case	1.9	3*5	3.4	3.7	4.5	3.5	3.2	4.8	3.5

eighth week, and in another case (1269), which developed measles, treatment lasted for six weeks only. Consequently the number of cases under treatment was in the first week twenty-six and in the eighth week twenty-four.

The average weekly number, over a period of eight weeks, of cases which had (1) parasitic febrile relapses was 17'0 per cent. of cases treated, (2) non-parasitic febrile attacks 2'0 per cent., (3) febrile paroxysms (parasitic and non-parasitic) 19'0 per cent., and (4) parasitic relapses (febrile and non-febrile) 46'3 per cent. (Tables V and VI).

After treatment. Of the twenty-five cases observed after cessation of treatment fifteen relapsed within the sixty-day observation period. Parasites re-appeared in one to thirty-three days and febrile relapses occurred in one to nineteen days after cessation of treatment. In one (1268) of the twenty-six cases the full course of treatment was not completed as the condition was not controlled. This case therefore should be added to the cases that relapsed, making the total failures sixteen (62 per cent.).

In two of the ten cases which did not relapse the post-treatment observation period was less than sixty days, viz., in Case 1246, twenty-three days and in Case 1255, forty days. Consequently the minimum number of relapses (those actually observed) was sixteen, and the possible maximum number (those actually observed + the number of cases not relapsing but lost sight of before expiration of the post-treatment observation period of sixty days) was eighteen, giving a minimum percentage of 62 and a maximum of 69 (Tables XIII and XVI).

CONCLUSION

Daily administration of arsenic, or of arsenic and strychnine, in small doses (minims 15), in combination with quinne (grains 5) is not more effective than the same dose of quinine alone.

TABLE V.

E. Summary of results of oral administration of Quinine HCl grains 5 daily for 8 weeks, in simple tertian malaria.

Week of Treatment	ıst	2nd	3rd	4th	5th	6th	7th	8th
Number of cases treated	26	26	26	26	26	26	25	24
Number of cases having parasitic febrile relapses	1	7	6	7	+	2	5	3
Number of cases having non-parasitic febrile attacks		0	1	0	1	0	0	2
Crand total of all febrile cases	1	7	. 7	7	5	2	5	5
Total number of parasitic febrile relapses	1	12	9	8	+	3	7	4
Total number of non-parasitic febrile attacks		0	1	0	í	0	2	+
Crand total of all febrile par- oxysms	1	12	10	8	5	3	7	8
Number of parasitic cases (febrile and non-febrile »	1 1	13	I 2	11	13	I 2	1 3	10
Fotal number of occasions parasites found	27	55	59	: 47	52	56	53	37

Table VI.

Analysis of Table V.

Week of Treatment	Ist	2nd	3rd	4th	5th	6th	7th	8th	Average per week
Percentage of parasitic febrile relapse cases per cases treated	3.8	26.0	23.1	26.0	15.4	7:7	2(1-1	12.5	17:0
Number of parasitic febrile relapses per parasitic febrile relapse case	1:4	1.7	1.2	1 - 1	1.~	115	1.1	1-3	1 • 3
Percentage of all febrile (parasitic and non- parasitic) cases per cases treated	3.8	26.9	26.9	26.9	10.2	7.7	2011	2018	19•0
Number of all febrile (parasitic and non- parasitic) paroxysms per febrile case	1.0	1.7	1.1	1 - 1	1.0	1.2	1:4	1.6	1:3
Percentage of all parasitic (febrile and nou- febrile) cases per cases treated	42.3	50.0	46.5	42.3	50.0	46.2	52·c	41.7	46.3
Number of occasions parasites found per parasitic case	2.5	1.5	5.0	4.3	4.0	+.7	1.1	3.7	4.0

GROUP III. Large doses of arsenic

In view of the inefficiency of the small dose of arsenic (minims 15 daily) we proceeded to ascertain whether better results were obtainable by the use of double the dose (minims 30 daily).

F. Liquor arsenicalis minims 30 daily,* with one or two periods of intermission, for eight weeks (Cases 1270-1283).

It was intended to give the arsenic according to the following plan: four weeks on treatment, one week off treatment, three weeks on treatment; but in only seven of fourteen cases was it found possible to adhere to this plan. In five cases it was found necessary, owing to the development of symptoms of arsenical poisoning, to interrupt treatment on two occasions, and in the remaining two cases owing to loss of sight it was not possible to continue treatment beyond the sixth week (Table VII). In all cases toxic symptoms disappeared shortly after leaving off treatment.

TABLE VII.

Showing mode of administration of arsenic in the various cases in Series F.

Number	Serial		Duration				
of cases	numbers	Weeks on	Weeks off	Weeks on	Weeks off	Weeks on	treatment
7	1270 to 1272 and 1277 to 1280	4	I	3			8 weeks
5	1273 to 1276 and 1281	3	1	2	1	1	8 weeks
2	1282 and 1283	3	1	2		•••	6 weeks

In one of the fourteen cases treatment was commenced during an apyrexial period; in the remaining thirteen cases the temperature fell to normal within ten days of the beginning of treatment.

In thirteen of the fourteen cases parasites disappeared from the cutaneous blood in from two to six days. In the remaining case (1278) parasites persisted practically throughout treatment (Table XVII).

^{*} Liq. arsenicalis M x Aq. chloroformi ad \$\overline{3}\$ i \$\overline{3}\$i. t.d.s.

RELAPSES

During treatment. In two (1282 and 1283) of the fourteen cases, owing to the development of severe symptoms of arsenical poisoning, it was found impossible to continue the treatment beyond the sixth week. Consequently the number of cases under treatment was in the first week fourteen and in the eighth week twelve. The average weekly number, over a period of eight weeks, of cases which had (1) parasitic febrile relapses was 3.8 per cent. of cases treated, (2) non-parasitic febrile attacks 3.5 per cent., (3) febrile paroxysms (parasitic and non-parasitic) 7.3 per cent., and (4) parasitic relapses (febrile and non-febrile) 28.1 per cent. (Tables VIII and IX).

After treatment. Eleven of the thirteen cases (85 per cent.), observed after cessation of treatment, relapsed within the sixty-day observation period. Parasites re-appeared in one to thirty-nine days and febrile relapses occurred in one to forty-two days after cessation of treatment (Tables XIII and XVII).

TABLE VIII.

F. Summary of results of oral administration of Liquor arsenicalis minims 30 daily, with one or two intermissions, for 8 weeks, in simple tertian malaria.

Week of Treatment	ıst	2nd	3rd	4th	5th	6th	7 t lı	Stlı
Number of cases treated	14	14	14	14	14	14	12	12
Number of cases having parasitic febrile relapses	0	0	1	0	0	2	0	1
Number of cases having non-parasitic febrile attacks	1	1	1	τ	0	0	0	0
Crand total of all febrile cases	1	1	2	1	0	2	0	1
Total number of parasitic febrile relapses	0	0	í	0	0	6	0	7
Total number of non-parasitic febrile attacks	2	3	I	+	0	0	0	0
Grand total of all febrile par- oxysms	2	3	2	4	0	6	0	1
Number of parasitic cases (febrile and uon-febrile)	9	3	2	I	2	+	+	5
Total number of occasions parasites found	15	S	3	7	9	. 14	14	2 1

TABLE IX.

Analysis of TABLE VIII.

Week of Treatment	Ist	2nd	3rd	4th	5th	6th	7th	8th	Average per weck
Percentage of parasitic febrile relapse cases per cases treated	0	0	7.1	0	0	14.3	0	8.3	3.8
Number of parasitic febrile relapses per parasitic febrile relapse case	0	0	1.0	0	0	3.0	0	1.0	0.6
Percentage of all febrile (parasitic and non- parasitic) cases per cases treated	7.1	7.1	14.3	7.1	0	-14-3	0	8.3	7•3
Number of all febrile (parasitic and non- parasitic) paroxysms per febrile case	2.0	3.0	1.0	4.0	0	3.0	0	1.0	1.75
Percentage of all parasitic (febrile and non-febrile) cases per cases treated		21.1	14.3	7.1	14.3	28.6	33.3	41.7	28•1
Number of occasions parasites found per parasitic case	1.7	2.7	1.5	7.0	4.2	3.2	3.2	4.2	3.6

G. The same as Treatment F with an injection of Quinine 2 HCl grains 15 intramuscularly on each of the first two days only (Cases 1284-1316).

The object of this series of observations was to determine whether, having caused the disappearance of parasites from the cutaneous blood and having controlled symptoms by two initial intramuscular injections of quinine, the daily administration of Liquor arsenicalis minims 30 would suffice to prevent relapses (parasitic and febrile). The arsenic was given according to the following plan: two weeks on treatment, one week off, two weeks on, one week off, two weeks on; all cases were able to tolerate this.

In five of the thirty-three cases treatment was commenced during an apyrexial period; in the remaining twenty-eight cases the temperature fell to normal within two days of the beginning of treatment.

In all cases parasites disappeared from the cutaneous blood in from one to four days (Table XVIII).

TABLE X.

G. Summary of results of administration of Liquor arsenicalis minims 30 daily, with two periods of intermission, for 8 weeks, + Quinine 2 HCl grains 15 intramuscularly on each of the first two days only, in simple tertian malaria.

Week of Treatment	ıst	2nd	3rd	4th	5th	6th	7th	8th
Number of cases treated	33	33	32	32	32	32	32	31
Number of cases having parasitic febrile relapses	0	0	0	0	I	1	4	I
Number of cases having non-parasitic febrile attacks	0	3	3	2	I	3	0	3
Grand total of all febrile cases	0	3	3	2	2	4	4	4
Total number of parasitic febrile relapses Total number of non-parasitic febrile attacks	0	0	0 7	0	4	1 5	5	3
Crand total of all febrile par- oxysms	0	3	7	3	7	6	5	6
Number of parasitic cases (febrile and non-febrile)	0	0	0	I	2	1	4	2
Total number of occasions parasites found	0	0	0	2	8	6	13	13

TABLE XI.

Analysis of Table X.

Week of Treatment	Ist	2nd	3rd	4th	5th	6th	7th	8th	Average per week
Percentage of parasitic febrile relapse cases per cases treated	0	0	0	0	3.1	3.1	12.5	3.5	2.7
Number of parasitic febrile relapses per parasitic febrile relapse case	0	0	0	O	1.0	1.0	1.0	3.0	1.1
Percentage of all febrile (parasitic and non- parasitic) cases per cases treated	0	9*4	9.7	6.2	6.2	12.5	12.5	12.0	8.7
Number of all febrile (parasitic and non- parasitic) paroxysms per febrile case	0	1.0	2.3	1.5	3.2	1.2	1.2	1.5	1.6
P ecentage of all parasitic (febrile and non-febrile) cases per cases treated	0	0	0	3.1	6.2	3.1	12.5	6.4	3.9
Number of occasions parasites found per parastic case	0	0	0	2.0	1.0	6.0	3.2	6.5	2.7

RELAPSES

During treatment. In one (1315) of the thirty-three cases, owing to the severity of the relapses and the grave clinical condition of the patient, it was found impossible to continue the treatment beyond the seventh week; in another case (1316) owing to intercurrent disease, treatment was given for two weeks only. Consequently the number of cases under treatment was in the first week thirty-three and in the eighth week thirty-one. The average weekly number, over a period of eight weeks, of cases which had (1) parasitic febrile relapses was 2.7 per cent. of cases treated, (2) non-parasitic febrile attacks 6.0 per cent., (3) febrile paroxysms (parasitic and non-parasitic) 8.7 per cent., and (4) parasitic relapses (febrile and non-febrile) 3.9 per cent. (Tables X and XI).

After treatment. Three of the thirty-one cases observed after cessation of treatment relapsed within the sixty-day observation period. Parasites re-appeared in one to fifteen days and febrile relapses occurred in one to nineteen days after cessation of treatment. In one (1315) of the thirty-three cases the full course of treatment was not completed as the condition was uncontrolled. This case therefore should be added to the cases that relapsed, making the total failures four (12.5 per cent.). (Tables XIII and XVIII.)

H. Control series. Quinine 2 HCl grains 15 intramuscularly on each of two consecutive days only (Cases 1317-1346).

In three of the thirty cases treatment was commenced during an apyrexial period; in the remaining twenty-seven cases the temperature fell to normal within three days of the beginning of treatment.

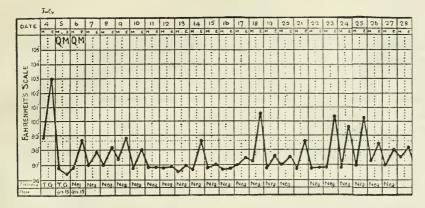
In all cases parasites disappeared from the cutaneous blood in from one to four days (Table $\rm XIX$).

RELAPSES

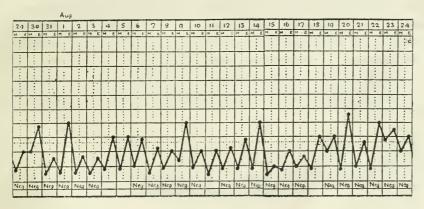
After treatment. Twenty-one of the thirty cases (70 per cent.) observed for sixty days after cessation of treatment relapsed. Parasites re-appeared in six to twenty-five days and febrile relapses occurred in nine to twenty-five days after cessation of treatment (Tables XIII and XIX).

389

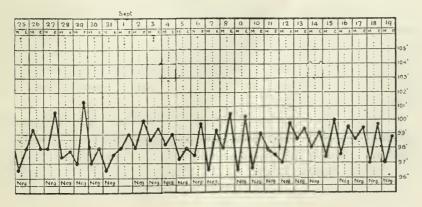
Case 1321 (Part I)



CASE 1321 (Part II)



CASE 1321 (Part III)



CONCLUSION

As a palliative, Liquor arsenicalis minims 30 daily appeared to exert a definite control of the temperature, the average weekly number of cases having parasitic febrile relapses being only 3.8 per cent. of cases treated; on the contrary, however, it failed in many cases to keep the cutaneous blood free from parasites, the average weekly number of cases having parasitic relapses (febrile and non-febrile) being 28.1 per cent. of cases treated. As a curative, the treatment was practically valueless, being followed by 85 per cent. of relapses.

The same treatment, however, with two initial injections of Quinine 2 HCl grains 15 gave very different results. The average weekly number of cases having parasitic febrile relapses was 2'7 per cent. of cases treated; but in contradistinction to the previous treatment the average weekly number of cases having parasitic relapses (febrile and non-febrile) was only 3'9 per cent. of cases treated. Furthermore, as a *curative* the result of this treatment (G) presented a striking contrast to that of the previous one (F), only 12'5 per cent. of cases relapsing within the post-treatment observation period of sixty days.

COMPARISON OF RESULTS OBTAINED FROM THE VARIOUS TREATMENTS

PALLIATIVE

From the figures relating to series C, D and E it will be seen that there is no evidence that small doses of arsenic (minims 15 daily) in combination with quinine are more efficacious than the same dose of quinine alone. There is also no evidence that a combination of small doses of arsenic and strychnine with quinine is more efficacious than quinine alone.

From the figures relating to F and G it will be seen that Liquor arsenicalis alone in large doses (minims 30 daily for eight weeks) effectively controls the febrile relapses; but if reliance were placed solely upon febrile attacks as indicating infection a fallacious

Table XII.

Comparison of palliative results obtained from the different treatments.

	С	D	E	F	G
		Liquor arsenici HCl minims 15 and Liquor strychninae HCl minims 15 + Quinine HCl grains 5 daily for 8 weeks	Quinine HCl grains 5 daily for 8 weeks	Liquor arsenicalis minims 30 daily	Liquor arsenicalis minims 30 daily with two periods of inter- mission for 8 weeks + Quinine 2 HCl grains 15 intramuscularly on each of the first two days only
Number of cases treated	28	25	26	14	33
Percentage of parasitic febrile relapse cases per cases treated (average per week)	25.7	13.2	17.0	3.8	2.7
Percentage of all febrile (parasitic and non-parasitic) cases per cases treated (average per week)	32.0	20.3	19.0	7:3	8.7
Percentage of all parasitic (febrile and non-febrile) cases per cases treated (average per week)	12:1	37.6	46.3	28-1	3.9

Table XIII.

Comparison of curative results obtained from the different treatments.

series	Treatment	Number of cases observed after	Number of cases which relapsed	Number of cases not relapsing but observed for less then 60	Percentage o	f cases which
		treatment*	within 60 days *	days	Minimum	Maximui
A	Liquor arsenicalis, minims 15 daily	3	3		100.0	100.0
В	Liquor arsenicalis minims 15 daily for 8 weeks + Quinine 2 HCl grains 15 intramuscularly, on each of the first two days only	2	2	***	100.0	100.0
С	Liquor arsenici HCl minims 15 + Quinine HCl grains 5 daily for 8 weeks	28	23	I	82:0	86.0
D	Liquor arsenici HCl minims 15 and Liquor strychninae HCl minims 15 + Quinine HCl grains 5 daily for 8 weeks	2.1	18	2	75.0	83.0
E	Quinine HCl grains 5 daily for 8 weeks	26	16	2	62.0	69.0
F	Liquor arsenicalis, minims 30 daily, with one or two periods of intermission, for 8 weeks	13	11	•••	85.0	85.0
G	Liquor arsenicalis, minims 30 daily, with two periods of intermission, for 8 weeks + Quinine 2 HCl grains 15 intramuscularly on each of the first two days only	3.2	+		12.2	12.5
н	Quinine 2 HCl grains 15 intramuscularly on each of two consecutive days only	30	21	•••	70.0	70.0

^{*} Including those cases in which treatment was abandoned as the condition was uncontrolled.

estimate of its efficacy would be obtained, for Table XII shows that whereas the figure in series F for parasitic febrile relapse cases is 3.8 per cent., that of all parasitic relapse cases, both febrile and nonfebrile, is 28.1 per cent. Arsenic in large doses, then, is anti-periodic in the sense that it controls the fever, but its control over parasites is much less. Further, the combination of quinine and arsenic in large doses is superior to arsenic alone, the total number of parasitic relapse cases being 3.9 per cent. for the combination and 28.1 per cent for arsenic alone. Expressed in clinical terms, the combination of large doses of arsenic and quinine controls the patients' fever and renders their blood negative during the course of treatment to an extent that arsenic alone is incapable of effecting.

CURATIVE

It will be seen in Table XIII that there is little to choose between any of the treatments with the exception of G, which is a combination of large doses of arsenic with quinine. This treatment gave only 12.5 per cent. of relapses within the sixty-day post treatment observation period.

TABLE NIV.

C. Results of oral administration of Liquor arsenici HCl minims 15 + Quinine HCl grains 5 daily for 8 weeks, in simple tertian malaria.

† E.A. = East Africa. M. = Mesopotamia. S. = Salonika.

Remarks				ioto E. on 5th, 100° E.	on 8th, 103 F. on 12th, 100 F. on 17th days; 1010 F. on 18th, 103 F. on 19th, 104:5 F. on 2+th	days.								Quinine orally grs. 30	on 11th day.
Observation period (in days) in cases	which did not	relapse	:	:		:	:	37	:	:	:	:	:	:	10
Parasitic (above Obserrellpse 100° E.) vation occurred occurred occurred in—days in—days in cases	cessation of	treat- ment	9	37		61	~	:	91	'n	4%	eţ	c1	:	:
Parasitic relapse occurred in—days after	cessation of	treat- ment	4	3437		13	-	:	15	П	ç	-	I	10	:
-		8th	1 + P3	0		2*	5+2	0	0	ď	-	Pi	3 + P2	Å	0
Isitic lood t		7th	1+P3	0		*	1 + P	0	0	3 + P3	1+P2	P	-	8	0
ms (parallts of b	nt	6th	0	*		*	0	0	0	1 + PI	1+P1	Pa	0	0	0
Number of febrile paroxysms (parasitic and non-parasitic) and results of blood examinations during treatment	Week of Ireatment	5th	. •	0		0	0	0	0	0	*	-i-	2+P1	0	0
febrile trasitic) ations d	eek of	†th	P3 1*	0		**	3 + P2	0	0	Pe 1*	**	Ā,	0	**	0
mber of non-pa examin	5	3rd		С		0	2+P2	0	0		*	0	0	0	*
Nun	-	puz	0	0	ಎಣಗ್ ಚ	0	P3 2+P5	٥ •	P 0	P2 2+P2	0	P 3	0	0	0
		18t	.	0		*	Д.	4		<u>µ</u>	0		С	0	0
	_	first dose	I-2	"		ες.	:	10	10	:	7	7	80	-	3
Temperature fell to normal in — days	after first dose		Same day	a		-	-	<i>w</i>	-	8	61	10	и	Same day	63
Date of end of	treatment		81.6.9	81.6.9		6.9.18	81.6.6	12.9.18	12.9.18	13.9.18	13.9.18	17.9.18	81.6.81	81.6.61	20.9.18
Interval (in months) between arrival in England	and	treatment	+	9		+	+	+	14,		+	+	- 1	15.	3
Interval Interval (in months) (in months) between leaving arrival in infected England	area and present	treatment	+	^		+	+	9	+	+	+	+	19	15	3
1 (84)	with	and present treatment	12	0		7	1.8	16	7	12	4	15	36	56	15
Place of in-	rection		si	ઝં		ઝં	×.	ś	ś	Š	ś	ŝ	E.A.	ŝ	S.
Number	case	4	1611	1192		1193	1611	1195	9611	1107	1198	6611	1200	1201	1202

‡ Three days after commencement of treatment are allowed for the disappearance of parasites and for the fall of the temperature to normal; consequently, the records for the first week refer to the last three days only of that week.

TABLE NIV-continued.

	Remarks			th days; 103.8° F.	on 7th day.		days: 100-t° F. on	19th day, 100° F. on 22nd day. Quinine orally grs. 30 on	15th day Pneumonia on cessation	or treatment.	Quinine orally grs. 30 on	Bronchial pneumonia on	Quinine orally grs. 30 on	Treatment changed: condition uncon-	trolled. Quinine intramuscularly in 8th week	Quinine intramuscularly	Quinine intramuscularly	Quinine intramuscularly in 5th week.
Observation period (in days)		redpos	:	:	89	÷	99	:	:	:	:	96	:	:	:	:	:	:
Febrile (above 100° F.) occurred in—days	after cessation of	treat- ment	10	11	:	-	:	:	17	1.3	:	:	:	:	:	:	:	:
Parasitic relapse occurred in—days after	cessation of treat-	ment	-	11-12	;	-	:	ż	81	Ξ	6	:	Ξ	:	:	:	:	:
		8th	P	*	0	$2 + P_5$	*	0	P	ď	ď	0	**	:	:	:	:	:
sitic		7th	Ps	0	0	3+P4	0	0	1 + P3	ď	1 + P2	0	D 2	7	2 + P3	:	:	:
Number of febrile paroxysms (parasitic and non-parasitic) and results of blood examinations during treatment	nt	6th	Pa	*	0	P,	0	0	4 + P3	м	-	0	*	7	ď	1 + P5	:	:
oaroxysu ind resu iring tro	Week of Treatment	5th	0	0	0	$1 + P_6$	0	*	Å	P	_	0	P	က	0	2+P5	:	:
febrile pasitic) a	eek of T	4th	0	0	0	2 + P5	0	0	1 + P6	P ₂	P	0	ď	Pı	0	4 + P2	2 + P3	2 + P5
non-par	W	3rd	0	0	0	2 + P5	0	0	P7	- P5	μ	0	*	1+P5	0	*	0	2 + P5
Num		2nd	0	0	0	2 + P:	0	0	2 + P4	Ę,	1 + P5	0	0	2+P5	0	0	C·	Pı
		ıst	0	Pı	0	1 + P2	떤	0	Ę.	P2	0	0	0	P2	0	ď	0	P
Parasites disappeared from	blood in — days	after first dose	77	10	eĭ	:	10	4	:	;	r1	eī	rì	:	+	9	C3	:
Tempera-	in — days after first dosc	hrst dosc	63	63	c)	ч	c1	درا	+	61	Apyrexia	1	Same day	w	c1	61	Apyrexia	**
Date	end of treatment		23.9.18	24.9.18	30.9.18	4.10.18	9.10.18	15.10.18	17.10.18	17.10.18	18.10.18	18.10.18	21.10.18	13.10.18	81.6.11	30.8.18	81.6.01	10.9.18
Interval (in months) between	England and present	present	+	9	¢1	61	*+	~	3	9	şv.	+	'n	:	m	'n	10	+
Interval Interval (in months) (in months) between between longer of the property of the proper	infected area and present	present treatment	+	12	61	+	10,	*+	ε,	7	ıv	9	4	:	*+	9	+	ıν
ths)	to a hospital with	malarra and present treatment	26	17	1.1	28	16.	61	56	1.3	27	28	0+	:	r.	11	r,	15
i d	of in- fection		s.	s:	s.	E.A.	s.	ઝ	s.	S.	ŝ	s.	M.	:	s.	sć.	s.	s.
	of		1203	1204	1205	1206	1207	1208	1209	1210	1211	1212	1213	1214	1215	1216	1217	1218

374

Results of oral administration of Liquor arsenici HCl minims 15 and Liquor strychninae HCl minims 15 + Quinine HCl grains 5 daily for 8 weeks, in simple tertian malaria. Ö.

† E.= Egypt. E.A.= East Africa. S.= Salonika.

Remarks			Not observed after	treatment. Quinine intramuscularly	on 1st day.			Quinine orally grs. 45	on 21st day.		100-8° F. on 35th day.	
			Not ol	treat Quinin	no no			Quinin	on 2		100.80	
Observation period (in days) in cases which did not relanse relanse	-	:	:	:	:	+2	:	:	:	38	69	:
Parasitic relapse relapse (above coeured) and after in—days cosation after of cessation frequency.	treat- ment	12	:	;	OI	:	S	:	+1	:	:	01
Parasitic relapse crelapse (above cocurred 100°); in—days cccurred after in—day ccssation after of cssation of creat of cocurred of creat of cocurred treat of cocurred cocurr	ment	6	:	-	01	:	7	-	-	:	:	w
	8th	0	0	P ₇	0	0	0	P	$1+P_6$	0	*	ď
sitic	7th	0	0	1 + Pe	0	0	Pı	Ā	å	0	0	0
Number of febrile paroxysms (parasitic and non-parasitic) and results of blood examinations during treatment Week of Treatment	6th	0	0	1 + P5	**	0	0	0	1 + P2	0	0	0
of febrile paroxysms varasitie) and results nations during treat	Şth	0	0	$2+\mathbf{P}_{2}$	0	0	0	å	m	0	0	0
ebrile Fasitic) a frions du	+th	0	0	å	ď	0	0	P.	1 + P2	0	0	С
ber of f non-par xaminal	3rd	0	0	ď	Ē,	مُ	0	O	0	0	0	0
Num and r	2nd	0	0	P	1 - P6	Pı	P	0	*	0	0	Ď,
	IST	0	0	+ + P3	က	0	*	ď	2	ď	0	ď
Parasites disappeared from cutaneous blood in — days after	first dose	-	¢1	15	:	+	ਜ	4	9	w	6	10
<u> </u>	first dose	Apyrexia 1	el	+	∞	-	7	ci ci	9 9	Same day 5	ct 50	10
Date ture fell from of to normal end of in — days bood treatment first dose	first dosc	7.9.18 Apyrexia 1		8.9.18	8.9.18	8.9.18	4 4 4					
Date ture fell of to normal end of in diversification freatment first dose	treatment first dose		r?	+	∞	4 8.9.18 1	3 9.0.18 + 4	et	9	Same day	ct	er.
Date ture fell of to normal end of in diversification freatment first dose			r?	8.9.18	8.9.18			11.9.18	12.9.18	12.9.18 Same day	17.9.18	17.0.18
ths) Interval (in months) (in months) between between between Date ture fell leaving arrival in of to normal infected England end of in edys area and and treatment first dose	treatment	+ 7.9.18	r?	8.9.18	6 8.9.18 8			11.9.18	12.9.18	8 12.9.18 Same day	17.9.18	17.0.18
(in months) (in months) between between between Date ture fell leaving arrival in of to normal infected England end of in days area and and treatment first dose present	treatment treatment	5 + 7.9.18	+ + 7.9.18	+ 3 8.9.18 +	8 6 8.9.18 8	+	+	+ 3 11.9.18 2	7 6 12.9.18 6	8 8 12.9.18 Same day	4 3 17.9.18 2	4 3 17.0.18

#Three days after commencement of treatment are allowed for the disappearance of parasites and for the fall of the temperature to normal; consequently, the records for the first week refer to the last three days only of that week.

Truck NV - continued.

	Remarks		100° F. on 3rd, 5th and			Collosol manganese on 2rd day.			Quinine orally grs. 45	Influenza in 7th, pneu-	Quinine orally grs. 30	Bronchial pneumonia on cessation of treatment.		Quinine orally grs. 15 on roth day	Quinine orally grs. 30 on	Quinine intramuscularly in 6th week.	
Observation period (in days) in cases	which did not relapse		:	:	99	:	:	:	:	63	:	77	:	:	:	:	
Febrile relapse (above 100° F.)	after cessation of	treat- ment	7	∞	:	:	7	27	:	:	:	:	10	:	:	:	
Parasitic relapse occurred in—days after	cessation of treat-	ment	Ξ	∞	:		∞	97	61	:	-	:	100	-	^	:	
		8th	#	0	0	P	0	0	0	2;	Pe	က	0	Ĺ.	P.	:	
sitic		7th	0	0	0	ď	0	0	0	*	ğ	PA	P	P	Ž,	:	
Number of febrile paroxysms (parasitic and non-parasitic) and results of blood examinations during treatment	nt	6th	1+10	*	0	Ps	ď	0	0	0	_	0	Ē,	0	ď.	:	
aroxysn nd resul iring tre	Week of Treatment	Sth	2 + P2	0	0	0	0	0	0	0	1+P4	0	**	2	P ₇	4+P3	
ebrile p asitic) a tions du	ek of T	4th	0	0	0	0	0	0	0	0	2 + P5	Pa	*	m	P. P.	4	
ber of f 10n-par xamina	We	3rd	-	0	0	0	*	0	0	0	Ž,	P7	*	be L	P	1 + P.	
Num and		puz	**	*	0	0	0	0	0	0	3 + P3	ğ	0	ú	1+P4	£ .	
		ıst	0	က	0	0	0	0	0	0	0	Ĕ,	0	-	ž	1+P1	
Parasites disappeared from	blood in — days after	inst dosc	+	7	eţ	+	61	es	_	+	et	7	u	:	:	:	
Fempera- ture fell	in — days after first dose	nrst dosc	10	7	ci	m	eş.	pet	pet	c1	pel	1	Apyrexia	Apyrexia	Apyrexia	e	
Date	end of treatment		20.9.18	20.9.18	22.9.18	24.9.18	27-9-18	4.10.18	9.10.18	18.10.18	18.10.18	23.10.18	24.10.18	24.10.18	28.10.18	11.9.18	
Interval (in months) between	England and	present treatment	+	\$	7	- 61	ıv.	10.	*†	m	6	9	rı	:	+	н	
Interval Interval (in months) (in months) between between Interval	infected area and	present	+	~	∞	8	7	W	10	ν.	6	7	ri	:	v.	7	
ll lhs)	hospital with	and present treatment	7.	1.2	7.7	13	15	15	Ξ	c 3	91	28	01	:	23	13	
ā	of in- fection		ઝં	ķ	sć.	si.	sć.	s.	si.	ಪ	ઝં	sć.	si.	ઝ	ý.	E.A.	
	of		1230	231	1232	1233	1234	1235	1236	1237	1238	1239	1240	1241	12:42	1243	

390

† During treatment.

E. Results of oral administration of Quinine HCl grains 5 daily for 8 weeks, in simple tertian malaria.

S. = Salonika.	
E.A. = East Africa.	
E.A. ==	
E Egypt.	
+	

-	Kemarks			397		(bronchitis); 100° F.	on 40th day.	Collosol manganese on 2nd day.	Quinine orally on 23rd day.		Collosol manganese on 2nd day.		
	which did not relapse		:	:	23	\$:	:	:	98 	:	:	70
Parasiuc relapse relapse (above occurred 100° F.) in—days occurred after in—days	Ö	treat- ment	×	61	:	:	:	:	:	:	:		:
Parasitic relapse relapse (above occurred 100° F.) in—days occurre after in—day	cessation of treat-	ment	-	-	:	:	5	1	12	:	-	el .	:
		Sth	ي	1 + P	0	*	å	P ₇	0	0		*	0
sitic		7th	P7 1 +P5	С	0	0	P1 2+P2	P	0	С	2 + P ₅	-	0
s (para s of blo tment	44	6th	p 7	PA	0	0	Pı	P	0	0	ř	<u>Q</u>	0
Number of febrile paroxysms (parasitic and non-parasitic) and results of blood examinations during treatment	Week of Treatment	şth	1+P4	P7	0	0	1 + P3	P7	Pı	0	P.	P4	*
brile pa itic) an	k of Tr	4th	+ Pu 4+P2 2+P3 1+P1 1+P1	$2+\mathbf{P}_{1}$	0	0	1+P4 1+P4 1+P3	Ъ	0	0	1 - P4	0	0
er of fe n-paras minatio	Wee	3rd	2 + P5	Pe	*	0	1 + P.	2 + P5	0	0	Pos	مُل	0
Numb and no exa		puz	+ P2	P3 1 + P6	0	0		P3 1+P5 2+P5	Pa	0	P	å	0
		1 St	+ Pa 4	P3.	0	Ō	0	P3	Ä	0	$1+\mathbf{p}_2$		0
Parasites disappeared from	blood in — days	first dose	:	:	ć	10	15	* * *	10	el	*	c1	·~
	in — days after fret dose	7007	cl	н	^1	-	rı	e)	-	-	Same day	-	_
Date	on end of treatment		81-6-1	7-9-18	81.6.6	12.9.18	15.9.18	25.9.18	25.9.18	25.9.18	25.9.18	25.9.18	26.9.18
Interval (in months) between	England and	present treatment	9	647	+		+	. 9	+	رى 	~	N	(2)
Interval Interval (in months) between	leaving infected area and	present treatment	9	10	-1	+	-1	. 0	v	+	+	10	+
Interval (in months) between first admission	to a hospital with	malaria and present treatment	55	. 6	12	ci	-	13	111	3	37	+	
	fPlace of in- fection		ś	E.A.	ŝ	ú	vi	જે :	s.	sć.	أتشأ	si.	si.
	Number of case		-	1 1 2 2 2	1216	1247	200	1240	1250	1251	1252	1253	1254

*Three days after commencement of treatment are allowed for the disappearance of parasites and for the fall of the temperature to normal; consequently, the records for the first week refer to the last three days only of that week.

TABLE XVI -continued.

				n		39 14	18							п	n		
	Remarks			Quinine orally grs. 30 on	Sym day.	ETS.	on 15th day.						No febrile relapse in	Quinine orally grs. 30 on	Quinine orally grs. 30 on	Treatment changed.	Measles in 1st week.
Obser- vation period (in days)			<u>o</u>	:	65	:	:	70	:	:	66	19	:	:	:	:	\$.
Febrile relapse (above 100° F.)	after cessation of	treat- ment	:	:	:	:	15	:	10	6	:	:	:	:	:	:	:
Parasitic relapse relapse (above occurred 100°F.) in—days occurred after in—days	cessation of treat-	ment	:	22	:	15	1.5	:	+	-	:	:	I	¢1	-	:	:
		8th	0	ā	0	0	$1 + P^3 + P^1$	c	P	Pe	0	0	0	0	$2 + P_1$:	:
itic ood		7th	0	ď	0	P	1 + P.	0	یْم	P ₇	0	0	0	ď	Pe	Å	:
Number of febrile paroxysms (parasitic and non-parasitic) and results of blood examinations during treatment	1t	6th	0	С	0	P ₇	0	0	Å	1 + Pe	С	c	ď	0	2 + P5	Ps	0
aroxysn nd resul iring tro	reatmer	th.	0	0	0	Ъe	0	0	-	Å	0	0	0	1 + P3	2	Pı	0
chrile pasitic) a	Week of Treatment	+th	0	0	0	Pe	1 + P ³	0	ď	1 + Pe 1 + P2	0	0	0	-	0	P6	0
ber of f non-par	We	3rd	0	0	0	P	-	0	Ρί		0	0	0	7	0	۵	0
Num		2nd	0	Г	0	2 + P4	0	0	1+793	Ğ.	0	0	0	3 2 + PI	0	<u>ب</u>	0
		181	0	0	ď.	Б	0	0	0	P	0	0	P	Pa	0	P3	0
Parasites disappeared from	blood in — days after	first dose	r1	c)	10	÷	2: 33	c1	eş.	:	er.	-	10.	÷	(C	÷	-
Tempera- ture fell	in — days after first dosc			-	N	en,	el	Same day	п	m	п	Same day	-	et	-	Same day	Same day
Date	end of treatment		26.9.18	26.9.18	27.9.18	27.9.18	81.6.62	1.10.18	13.10.18	14.10.18	14.10.18	18.10.18	21.10.18	21.10.18	22.10.18	22.0.18	17.9.18
Interval (in months) between	England and present	treatment	61	-+	10	7	ro	7	ı,	7	-+	10	+	1/3	10	m	+
Interval Interval (in months) (in wonths) between between between between in trivial in	infected area and present	treatment	rc.	9	9	×	+	×	10.	01	+	10	20	10	10	-+	+
· ·	hospital with malaria	and present treatment	7.7	†1	23	21	13	% *1	27	91	7.	28	9	28	15	27	41
D	of in- fection		જ	s.	si.	ŝ	Š	જં	s;	EA.	s;	ઝં	s.	s.	တ်	s,	s;
Z Sub-	of case		1255	1256	1257	1258	1259	1260	1971	1262	1263	1971	1265	1266	1267	1268	1269

F. Results of oral administration of Liquor arsenicalis minims 30 daily, with one or two intermissions. for 8 weeks, in simple tertian malaria.

+ E.A. = East Africa. S. = Salonika.

						3	99										
	Remarks		(post-treatment)	Influenza first 2 weeks.	100.4° E. on 4th, 100° E. on 58th, 61st and 66th days.		Not observed.					No febrile relapse in 44	•		Symptoms of arsenical poisonings. Discharged	on 8th day, Treatment stopped owing to total but temporary blindness.	No febrile relapse in 62 days but parasites present daily.
Observation period in days)	which did not relapse		:	:	10.3	99	:	:	:	:	:	:	:	:	:	:	
Febrile relapse (above 100° F.) occurred (in days		treat- ment	67	91	-	:	:	-	. 11		×	:	01	39	:	:	
Parasitic relapse occurred in—days of after it	-	ment	=	11	:	;	:	1	66	1.5	ın.	13	1	39	os.	∞	
0.5		Sth	Ę,	Ç	0	0	0	1 + P5	0	0	Pe	P	4	0	:	:	
ood ood		7th	ď	0	0 .	0	0	0	0	Pa	P	0	D.	0	:	:	
nber of febrile paroxysms (paras non-parasitic) and results of blo examinations during treatment	nt	6th	0	10	0	0		С	0	P	5 + P	С	1 P5		<u>۾</u>		
oaroxysn ind resu uring tre	Week of Treatment	şth	0	0	0	0	0	0		0	7 Po	0	P	0	0		
febrile prasitic) ations d	eck of 1	+th	0		o 	0		0	٠	0	Pt P7		0	o 	С	* *	
Number of febrile paroxysms (parasitic and non-parasitic) and results of blood examinations during treatment	=	3rd	0	0		С	0		Pt 1+Pt		P. I	0			Б а	*	
Nu and		2nd	0	*	P 1	0	P ₁	Ре	P	0	P3	-	0	P ₁	ا ا	A	
	1	1st	0	*2*		0				0		0					
Parasites disappeared from	blood in — days	first dose	10	10	16,	ы	16	ي	9	ces	:	+	w	çe,	4	9	
	in — days after first dosc		Sanic day	01	-1	Same day	r1	+	ı	v.	÷		ee.	Apyrexia	c1	er,	
Date of	end of treatment		21.8.18	24.8.18	24.8.18	26.8.18	26.8.18	27.8.18	28.8.18	28.8.18	20,8.18	4.0.18	16.9.18	28.9.18	30.7.18	10.8.18	
Interval in months) between	England and	treatment	(1)	61	(e)	rl	ev.	10	r)	7	11	rc.	15	9	tol (C1	
Interval Interval (in months) (in months) between between Interval in months)	infected area and	٠,		9	(C)	٠٠,	ec	50,	55,	1	V	(C	÷	×	***	el	
al ths)	to a hospital with		12	6	1 <i>2</i> °,	56	ir -	10	. CI	11	1,	77	ci ci	2.1	er,	±	
	of in- fection		si.	F1.	si.	ś	νί.	×	ઝં	vá.	ઝં	s.	÷.	E.A.	vi.	sć.	
	of case		1278	1271	61 61 61	2,2	1274	14	1276	17	1278	0221	128	1281	1282	× 1	

Three days after commencement of treatment are allowed for the disappearance of parasites and for the fall of the temperature to normal: consequently, the records for the first week refer to the last three days only of that week. § During treatment.

G. Results of administration of Liquor arsenicalis minims 30 daily, with two periods of intermission, for 8 weeks + Quinine 2 HCl grains 15 intramuscularly on each of the first two days only.

† E.= Egypt. E.A. = East Africa. S. = Salonika.

			arsenical			100	,.			ıy.	Jo p		th1 rrhal	.y.	рı		.;.	
	Remarks			-	n 33rd d		ı 55th day			n 13th da	ved: die	na.	emperature in 4th1 week due to catarrhal	n roth da	n 15th ai	į	n 23rd da	
	~		Symptoms of	poteod	100° F. on 33rd day		100° F. on 55th day.			102° F. on 13th day.	Not observed: died of	pucamouna.	Temperature in 4th1	cold.	100° F. on 15th and		100° F. on 23rd day.	
Observation period (in days			65	71	69	137	(i.7	:	47.	69	:	:	7.4	20	20	63	+9	
Parasitic relapse relapse (above occurred 100° F.) in—days occurred after in—days		treat- ment	:	:	:	:	:	61	:	:	:	-	:	:	:	:	:	
Parasitic relapse occurred in—days	cessation of treat-	ment	:	:	:	:	:	н	:	:	:		:	:	:	:	:	
		8th	٥	0	0	0	0	Pe	0	0	0	3 + P ₄	0	O	0	0	*	
asitic blood nt		7th	0	0	0	0	0	1 + P.	0	0	0	-	0	0	0	0	0	
Number of febrile paroxysms (parasitic and non-parasitic) and results of blood examinations during treatment	cnt	eth	0	0	0	0	0	3 1 + P5	0	0	0	0	0	0	0	0	0	Andrew or the second
paroxy and res	Week of Treatment	5 th	0	0	0	ď	0	c 4 + P3	0	0	0	0	0	0	0	0	0	1
febrilc arasitic) nations	reck of	+11	0	0	0	0	0	ğ	0	0	0	0	**	0	0	0	0	
mber of non-p exami	>	3rd	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	eatment
Nar		200	0	0	0	0	0	*	0	0	0	0	υ	0	0	0	0	1 During treatment.
		ıst	0	0	0	0	0	0	0	0	0	0	c	0	0	0	0	† Dr
Parasites disappeared from	blood ın — days alter	first dose	ci.	-	eı	-	61	_	-	61	r)	+	-	m	61	cı	63	
Tempera- ture fell	in — days after first dose		-	-	-	5.9.18 Apyrexia	-	-	14.9.18 Apyrexia	Same day	Same day	-	ы	ы	Ħ	Apyrevia) and	
Date	end of treatment		4.9.18	5.9.18	5.9.18	81.6.5	8.9.18	12.9.18	81.6.41	17.9.18	81.6.61	20.9.18	20.9.18	23.9.18	24.9.18	\$1.9.18	26.9.18	
Interval (in months) between	England and present	treatment	۲۲.	+	+	ee.	+	1	+	10	60	m	m	m	(A)	m	4	
Interval (in months) (in months) between between	infected area and present	treatment	"	+	+	+	v.	oc.	+	7	12	+		6	+	+	*+	A CONTRACTOR AND A CONT
(S)	hospital with malaria	and present treatment	23	7.7	+1	10.	+	91	r)	6	321	01	57	27	22	91	91	
——————————————————————————————————————	of in- fection		s.	s,	s,	s.	S.	ŝ	s.	E.A.	Š	ŝ	νż	s,	Š	Š	s;	
Simple			1284	1385	1286	1287	1288	1289	1290	1291	1292	1293	1671	5621	1296	1297	1298	

Number	Place	Interval (in months) between first admission to a		Interval Interval (in months) (in months) between leaving arrival in	Date	Tempera- cture fell to normal	Parasites disappeared from cutancous		Numl and n	Number of febrile paroxysms (parasitic and non-parasitic) and results of blood examinations during treatment	brile pa sitic) an ons dur	roxysm d result ing trea	s (parasi s of blo	fic	0.0	Parasitic relapse occurred in—days	Febrilc relapse (above 100° F.)	Observation period (in days)	
of	of in- fection	hospital with malaria	infected area and present	England and present	ent	in — days after first dosc	blood in — days after			Wee	k of Tr	Week of Treatment				=	after cessation of	which did not	Remarks
		and present treatment	treatment	treatment			first dose	ıst	puz	3rd	4th	th.	6th	7th	Sth.	ment	treat- ment		
1299	s,	23 H	+	-	31.9.18	-	c1	0	0	0	0	0	0	0	**	:	:	100	
1300	E.A.	13	6	+	2.10.18	H	61	0	0	*	0	0	0	0	0	:	:	117	101.6° F. on 12th day
1301	ŒĴ	91	V	W.	10.10.18	Same day		0	0	0	0	0	0	0	0	:	:	49	(bronchial catarrh).
1302	s.	SS 1881	IC	ਚ:	13.10.18	н	61	0	0	С	0	0	0	0	0	:	:	69	
1303	E.A.	120	~	9	81.01.61	_	-	0	0	*	0	0	0	0	0	:	:	92	
1304	si.	7.7	9	-+	16.10.18		N	0	C	0	0	*	0	0	0	:	:	104	
1305	ૹ૾	15	r)	c1	21.10.18	-	(0)	0	0	0	0	0	0	0	0	:	:	125	
1306	s.	1/r,	10.	10.	22.10.18	1	e)	0	0	0	*	c	2*	0		:	:	86	
1307	s;	n	+	+	22.10.18	1	ro.	0	0	, K	0	0	0	0	0	:	:	9	Influenza in 3rd week.1
1308	si.	00 61	01	5	23.10.18	Apyrexia	int.	0		0	0	0	0	0	0	:	:	96	
1300)	s.	<u></u>	ce	c1	23.10.18	Apyrexia	~	0	0	0	0	0	*	0	0	:	:	112	
1310	si.	91	en	61	24.10.18	r1	۲1	0	0	0	0	0	0	0		:	:	99	
1311	š	91	U°	v	24.10.18	-	^1	0	C	0	С	0	0	0	0		:	ود	
1312	×.	91	-+	m	25.10.18	tent	(n)	0	0	0	0	0	0	0	0	:		+6	
1313	E.A.	00	*+	c1	26.10.18	C1	en.	0	0	0	0	0	**	2	**	IC;	91	:	Influenza in 6th and
1314	š	35	es	61	24.8.18	Same day	n)	0	*	0	0	0	0	0	0	:	:	93	7th weeks.1 Symptoms of arsenical
1315	E.A.	0	7	en.	27.9.18	***	15	0	*	С	0	0	0	1 + P3	:	:	:	:	poisoning.1 Treatment changed.
1316	E.A.	1~	15	-	8.9.18	-	r1	О		:	:		:	:	:		:	09	104° F. on 9th day.

During treatment.

TABLE XIX.

H. Results of administration of Quinine 2 HCl grains 15 intramuscularly on each of two consecutive days only, in simple tertian malari

* E.A. = East Africa. F. = France. It. = Italy. S. = Salonika.

				(1	1		1
Number of case	*Place of in- fection	Interval (in months), between first admission to a hospital with malaria and present treatment	Interval	Interval (in months) between arrival in England and present treatment	Date of end of treatment	Temperature fell to normal in — days after first dose	Parasites disappeared from cutaneous blood in — days after first dose	Parasitic relapse occurred in — days after last dose	Febrile relapse (above 100°F.) occurred in — days after last dose	Observa- tion period (in days) in cases which did not relapse	Remarks
1317	S.	11	2	1	25.6.18	Apyrexia	2	18	18		
1318	S.	1 3	5	4	25.6.18	1	3	15	21	•••	
1319	s.	9	3	2	25.6.18	1	4	2.2	2.2		1
1320	S.	12	3	2	5.7.18	1	2	15	18		100°F. on 8th da
1321	S.	23	2	2	6.7.18	Same day	1			87	Irregular temperar after treatment Vide chart.
1322	S.	22	2	1	6.7.18	Same day	1	12	15		- 41
1323	s.	τ	2	1	6.7.18	3	1	23	24		
1324	s.	9	2	2	7.7.18	Same day	1	11	13	•••	
1325	s.	9	3	2	9.7.18	Same day	2	19	2.2	•••	
1326	S.	27	10	1	11.7.18	1	1	1.4	22	•••	
1327	. S.	23	10	1	12.7.18	1	1	18	2.2	•••	1
1328	S.	16	10	τ	12.7.18	I	2	1.2	13	•••	
1329	S.	25	2	2	14.7.18	• •••	2	25	•••	•••	
1330	It.	13	0	0	22.7.18	1	2	10	1.4	•••	
1331	s.	26	9	2	23.7.18	I	2	17	19		
1332	S.	11	2	I	25.7.18	Apyrexia	2		• •••	92	
1333	S.	10	2	I	27.7.18	Apyrexia	2			66	
1334	S.	11	7	6	27.7.18	1	2			73	101°E, on 24th day
1335	s.	12	4	3	28.7.18	Same day	2.	<i>;</i> ···		73	
1336	s.	23	4	2	28.7.18	Same day	1		•••	68	100°F. on 34th day
1337	E.A.	16	7	. 4	28.7.18	2	3	13	15		
1338	S.	21	6	5	28.7.18	2	2	20	21		
1339	S.	9	5	4	28.7.18	1	1			65	
1340	S.	15	6	5	28.7.18	I	2	16	15		
1341	E.A.	8	5	1	31.7.18	Same day	3	6	9	•••	
1342	F.	2	2	2,	8.8.18	Same day	3	23	25	•••	
1343	S.				14.8.18	1	3			71	
1344	S.	2.‡	3	3	22.8.18	I	1	21	24		
1345	S.	12	2	1	23.8.18	I	2	20	22		
1346	S.				23.8.18	ı	ī		***	62	100°F. on 18th, 24t and 25th days.